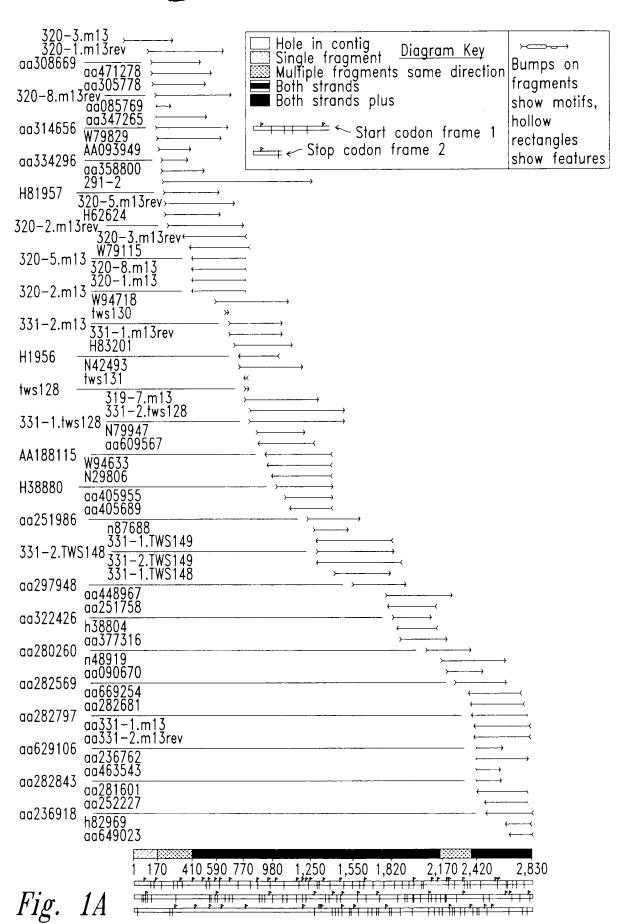
Inventor(s): Todd W. See Express Mail No. EL755724263US Docket PP-01406.004/200130.438D1



Inventor(s): Todd W. See

GAAGCAAGGAGGCGGCGGCGGCCGAGCGAGTGGCGAGTAGTGGAAACGTTGC TTCTGAGGGGAGCCCAAGATGACCGGTTCTAACGAGTTCAAGCTGAACCAGCC ACCCGAGGATGGCATCTCCTCCGTGAAGTTCAGCCCCAACACCTCCCAGTTCCT GCTTGTCTCCTCCTGGGACACGTCCGTGCGTCTCTACGATGTGCCGGCCAACTCC ATGCGGCTCAAGTACCAGCACACCGGCGCGTCCTGGACTGCGCCTTCTACGAT CCAACGCATGCCTGGAGTGGAGGACTAGATCATCAATTGAAAATGCATGATT TGAACACTGATCAAGAAAATCTTGTTGGGACCCATGATGCCCCCTATCAGATGT GTTGAATACTGTCCAGAAGTGAATGTGATGGTCACTGGAAGTTGGGATCAGA CAGTTAAACTGTGGGATCCCAGAACTCCTTGTAATGCTGGGACCTTCTCTCAGC CTGAAAAGGTATATACCCTCTCAGTGTCTGGAGACCGGCTGATTGTGGGAACA GCAGGCCGCAGAGTGTTGGTGTGGGACTTACGGAACATGGGTTACGTGCAGCA GCGCAGGGAGTCCAGCCTGAAATACCAGACTCGCTGCATACGAGCGTTTCCAA ACAAGCAGGGTTATGTATTAAGCTCTATTGAAGGCCGAGTGGCAGTTGAGTA TTTGGACCCAAGCCCTGAGGTACAGAAGAAGAAGTATGCCTTCAAATGTCAC AGACTAAAAGAAATAATATTGAGCAGATTTACCCAGTCAATGCCATTTCTT TTCACAATATCCACAATACATTTGCCACAGGTGGTTCTGATGGCTTTGTAAAT ATTTGGGATCCATTTAACAAAAAGCGACTGTGCCAATTCCATCGGTACCCCAC GAGCATCGCATCACTTGCCTTCAGTAATGATGGGACTACGCTTGCAATAGCGT CATCATATATGTATGAAATGGATGACACAGAACATCCTGAAGATGGTATCTT CATTCGCCAAGTGACAGATGCAGAAACCAAAACCCAAGTCACCATGTACTTGA CAAGATTTCATTTACTTAAGTGCCATGTTGATGATAATAAAACAATTCGTAC TCCCCAATGGTGGATTTATTACTATTAAAGAAACCAGGGAAAATATTAATTT TAATATTATAACAACCTGAAAATAATGGAAAAGAGGTTTTTGAATTTTTTT TTTAAATAAACACCTTCTTAAGTGCATGAGATGGTTTGATGGTTTGCTGCATT AAAGGTATTTGGGCAAACAAAATTGGAGGGCAAGTGACTGCAGTTTTGAGA AATAAGTGTAATAAAAATCCCTTTGCATTCTTTCTGGACCTTAAATGGTAGA GGAAAAGGCTCGTGAGCCATTTGTTTCTTTTGCTGGTTATAGTTGCTAATTCTA AAGCTGCTTCAGACTGCTTCATGAGGAGGTTAATCTACAATTAAACAATATT TCCTCTTGGCCGTCCATTATTTTCTGAAGCAGATGGTTCATCATTTCCTGGGCTG TTAAACAAAGCGAGGTTAAGGTTAGACTCTTGGGAATCAGCTAGTTTTCAAT CTTATTAGGGTGCAGAAGGAAAACTAATAAGAAAACCTCCTAATATCATTTT GTGACTGTAAACAATTATTTATTAGCAAACAATTGATCCCAGAAGGGCAAAT TGTTTGAGTCAGTAATGAGCTGAGAAAAGACAGAGCATATCTGTGTATTTGG AAAAATAATTGTAACGTAATTGCAGTGCATTTAGACAGGCATCTATTTGGAC CTGTTTCTATCTCTAAATGAATTTTTGGAAACATTAATGAGGTTTACATATTT CTCTGACATTTATATAGTTCTTATGTCCATTTCAGTTGACCAGCCGCTGGTGAT TAAAGTTAAAAAGAAAAAATTATAGTGAGAATGAGATTCATTTCAATGTA ATGCACTAAAGCAGAACACGAACTTAGCTTGGCCTATTCTAGGTAGTTCCAA CTCTGAGTAGGACTTGGACCTTTCCTGAGATTTATTTTATCCGTGATGTATTTT TITTAATTCTTTGATACAGAGAGGGTCTTTTTTTTTTTAAGTATTTCAGTGA AAACTTGGTGTAAGTCTGAACCCATCTTTTGAAATGTATTTTCTTCATTGCAG

Fig. 1B (Continued)

Alignment of predicted huBUB3 gene products from ~1.4 Kb cDNA and ~2.7 Kb cDNA

		20	40	60
huBUB3 2.7	MTGSNEFKLNQPPEDGI	SSVKFSPNTSQFLLVSSWD	TSVRLYDVPANSMRLKYQHT	ΓGAVL
huBUB3 1.4	MTGSNEFKLNQPPEDGI	SSVKFSPNTSQFLLVSSWD	TSVRLYDVPANSMRLKYQHT	rgavl
		80	100	120
huBUB3 2.7	DCAFYDPTHAWSGGLDH	IQLKMHDLNTDQENLVGTHD	APIRCVEYCPEVNVMVTGSV	VDQTV
huBUB3 1.4	DCAFYDPTHAWSGGLDH	IQLKMHDLNTDQENLVGTHD	APIRCVEYCPEVNVMVTGSV	VDQTV
				400
		140	160	180
huBUB3 2.7			RRVLVWDLRNMGYVQQRRES	
huBUB3 1.4	KLWDPRTPCNAGTFSQF	'EKVYTLSVSGDRLIVGTAG	RRVLVWDLRNMGYVQQRRES	SSLKY
		000	200	240
1 01100 0 7	OTDOID A EDNIKOOVIII O	200	220	240
huBUB3 2.7	•	·	KKYAFKCHRLKENNIEQIYF	
huBUB3 1.4	QTRCTRAFPNKQGYVLS	STEGRVAVEYLDPSPEVQK	KKYAFKCHRLKENNIEQIYF	NAI
		260	280	300
huBUB3 2.7	CEUNTUNTEATOCCDOE		TSIASLAFSNDGTTLAIASS	
		·	TSIASLAFSNDGTTLAIASS	
huBUB3 1.4	2 LUNTUM LA LAASTAL	VIVIWUPFINNNKLUQFINTF	131A3EAL3MAGLIFATA3) IIII L
		320		
huBUB3 2.7	MDDTEHPEDGIFIRQVT			
huBUB3 1.4	MDDTEHPEDGIFIRQVT			
	Inde			

1 -----MQIVQIEQAPKDYISDIKIIP---SKSLLL

27

scBUB3

1 -----SNEFKLNQPPEDGISSVKFSPN--TSQFLL 31 muBUB3 1 -----SNEFKLNOPPEDGISSVKFSPN--TSOFLL 31 huBUB3 58 1 MSLFGTTSGFGTSGTSMFGSATT--DNHNPMKDIEVTSSPDDSIGCLSFSPPTLPGNFLI rael-hu 46 1 MSLFG-----QATTSTVSNATG-----DLKKDVEVAQPPEDSISDLAFSP---QAEYLA rael 1 MSFFNR----SNTTSALGTSTAMANEKDLANDIVINSPAEDSISDIAFSP---QQDFMF 52 yet7 DI. 83 scBUB3 -ITSWDGSLTVYKFDIQAKNVDLLQSLRYKH--PLLCCNFI-DNTDLQIYVGTVQGEILK 81 muBUB3 -VSSWDT--SVRLYDVPA--NSM-R-LKYQHTGAVLDCAFY-DPT--HAWSGGLDHQLKM huBUB3 32 - VSSWDT--SVRLYDVPA--NSM-R-LKYQHTGAVLDCAFY-DPT--HAWSGGLDHQLKM 81 rael-hu 59 -AGSWAN--DVRCWEVQDSGQTIPK-AQQMHTGPVLDVCWSDDGS--KVFTASCDKTAKM 112 100 rae1 -ASSWDS--KVRIYEVOATGOSIGK-ALYEHOGPVLSVNWSRDGT--KVASGSVDKSAKV SASSWDG--KVRIWDVQN-GVPQGR-AQHESSSPVLCTRWSNDGT--KVASGGCDNALKL 106 yet7 SW .L D... scBUB3 VDLIGSPSFQALTNNEANLGICRICKYGD--DKLIAASWDGLIEVIDPRNYGDGVIAVKN 141 135 muBUB3 82 HDLNTDOENLVGTHDAPIRCVEYCPEVNV----MVTGSWDOTVKLWDPR-TPCNAGTFS-82 HOLNTDOENLYGTHDAPIRCVEYCPEVNV----MVTGSWDOTVKLWDPR-TPCNAGTFS-135 huBUB3 113 WDLSSNQAIQIAQHDAPVKTIHWIKAPNY--SCVMTGSWDKTLKFWDTR-SSNPMMVLQ-168 rael-hu rael 101 FDIQTGQNQQVAAHDDAVRCVRFVEAMGT-SPILATGSWDKTLKYWDLR-QSTPIATVS-157 164 YDIASGOTOQIGMHSAPIKVLRFVOCGPSNTECIVTGSWDKTIKYWDMR-QPQPVSTVMyet7 107 . . SWD . D. LNSNNTKVKNKIFTMDTNSSRLIVGMNNSQVQWFRLPLCEDDNGTIEESGLKYQIRDVAL scBUB3 201 muBUB3 136 Q-----PEKVYTLSVSGDRLIVGTAGRRVLVWDLWNMGYVQQRRE-SSLKYQTRCIRA 187 huBUB3 136 Q-----PEKVYTLSVSGDRLIVGTAGRRVLNWDLRNMGYVQQRRE-SSLKYQTRCIRA 187 rael-hu 169 L-----PERCYCADVIYPMAVVATAERGLIVYQLENQPSEFRRIE-SPLKHQHRCVAI 220 rae1 158 L-----PERVYAMDCVHPLLTVATAERNICVINLSEPTKIFKLAM-SPLKFQTRSLAC 209 165 M-----PERVYSMDNKQSLLVVATAERHIAIINLANPTTIFKATT-SPLKWQTRCVAC 216 yet7 . L SLKQR.

		 \/	
scBUB3 muBUB3 huBUB3 rae1-hu rae1 yet7	202 188 188 221 210 217	LPKEQEGYACSSIDGRVAVEFFDDQGDDYNSSKRFAFRCHRLNLKD-TNL FPNK-QGYVLSSIEGRVAVEYLDPS-PEV-QKKKYAFKCHRLKENN-IEQ FPNK-QGYVLSSIEGRVAVEYLDPS-PEV-QKKKYAFKCHRLKENN-IEQ FKDKQNKPTGFALGSIEGRVAIHYINPP-NPAKDNFTFKCHRSNGTNTSAPQD FIKGDGYAIGSVEGRCAIQNIDEKNASQNFSFRCHRNQAGN-SAD YNEADGYAIGSVEGRCSIRYIDDG-MQKKSGFSFKCHRQTNPN-RAPGSNGQSL G. SGRF.CHR	250 233 233 272 253 268
scBUB3 muBUB3 huBUB3 rael-hu rael yet7	251 234 234 273 254 269	AYPVNSIEFSPRHKFLYTÄGSDGIISCWNLQTRKKIKNFAKFNED-SVVKIACSDNILCL IYPVNAISFHNIHNTFATGGSDGFVNIWDPFNKKRLCQFHRYPTSIASLAFSNDGTTLAI IYPVNAISFHNIHNTFATGGSDGFVNIWDPFNKKRLCQFHRYPTSIASLAFSNDGTTLAI IYAVNGIAFHPVHGTLATVGSDGRFSFWDKDARTKLKTSEQLDQPISACCFNHNGNIFAY VYSVNSIAFHPQYGTFSTÄGSDGTFSFWDKDSHQRLKSYPNVGGTISCSTFNRTGDIFAY VYPVNSIAFHPLYGTFVTÄGGDGTFNFWDKNQRHRLKGYPTLQASIPVCSFNRNGSVFAY Y VN I F T G DG W	309 293 293 332 313 328
		TVN II I U DU W	
scBUB3 muBUB3 huBUB3 rae1-hu rae1 yet7	310 294 294 333 314 329	ATSDDTFKTNAAIDQTIELNASSIYIIFDYEN 341 ASSYMYEMDDT-EHPEDGIFIRQVTDAETKPKST 326 ASSYMYEMDDT-EHPEDGIFIRQVTDAETKPKSPC-T 328 ASSYDWSKGHEFYNPQKK-NYIFLRNA-AEELKPRNKK 368 AISYDWSKGYTFNNAQLP-NKIMLHPVPQDEIKPRPKKGR 352 ALSYDWHQGHMGNRPDYP-NVIRLHATTDEEVKEK-KK-R 365 A S E	

Fig. 3B

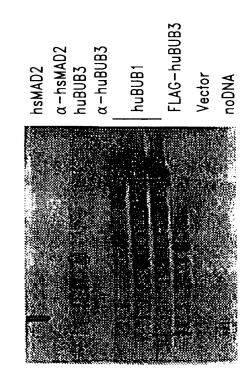


Fig. 4A

FLAG-huBUB3 huBUB3 + huBUB1 + + + hsMAD2 + SPSP α -FLAG SP S P huBUB1 (~121Kd) Fig. 4B FLAG-huBUB3 huBUB3 (~37Kd) hsMAD2 (23Kd)

mRNA expression of BUB/MAD homologs in various tissues.

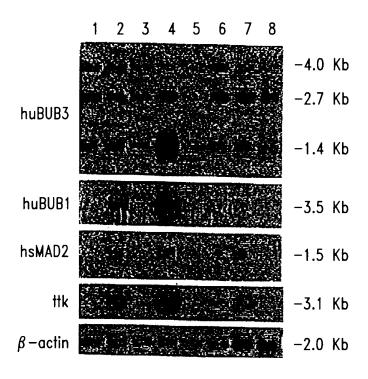


Fig. 5